# (19) World Intellectual Property Organization International Bureau



### | 1881 | 1887 | 1888 | 1881 | 1881 | 1884 | 1884 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1885 | 1

(43) International Publication Date 2 December 2004 (02.12.2004)

**PCT** 

## (10) International Publication Number WO 2004/103961 A3

(51) International Patent Classification<sup>7</sup>: C07D 217/04, A61K 31/47

(21) International Application Number:

PCT/US2004/015731

(22) International Filing Date: 19 May 2004 (19.05.2004)

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data:

60/471,346

19 May 2003 (19.05.2003) US

(71) Applicant (for all designated States except US): VIR-GINIA COMMONWEALTH UNIVERSITY [US/US]; 800 E. Leigh Street, Suite 113, Richmond, VA 23219 (US).

(72) Inventors; and

(75) Inventors/Applicants (for US only): DESAI, Umesh, R. [US/US]; 10190 Durhams Ferry Place, Mechanicsville, VA 23116 (US). GUNNARSSON, Gunnar [US/US]; 410 N. 12th Street #542, Richmond, VA 23298 (US).

(74) Agents: WHITHAM, Michael, E. et al.; 11491 Sunset Hills Road, Suite 340, Reston, VA 20190 (US).

(81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.

(84) Designated States (unless otherwise indicated, for every kind of regional protection available): ARIPO (BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

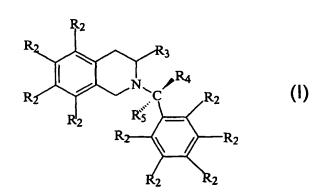
#### Published:

with international search report

(88) Date of publication of the international search report: 14 April 2005

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: SULFATED BIS-CYCLIC AGENTS



(57) Abstract: Sulfated bis-cyclic compounds that are potent anticoagulants and methods for their manufacture are provided. The sulfated compounds are bis-cyclic moieties comprised o an ioquinoline ring joined to a phenyl ring with the general chemical structure formula (I) where  $R_2$  is hydrogen or a sulfate moiety,  $R_3$  is either hydrogen or carboxylate moiety;  $R_4$  is a hydrogen or oxygen; and  $R_5$  is a hydrogen if  $R_4$  is a hydrogen, and is absent if  $R_4$  is oxygen. Counterions such as sodium may also be coordinated to the sulfate and carboxylate moieties.

WO 2004/103961 A3

### INTERNATIONAL SEARCH REPORT

International application No.

PCT/US04/15731

A. CLASSIFICATION OF SUBJECT MATTER				
IPC(7) : C07D 217/04; A61K 31/47				
US CL : 546/146,149; 514/307				
According to International Patent Classification (IPC) or to both national classification and IPC				
B. FIELDS SEARCHED				
Minimum documentation searched (classification system followed by classification symbols) U.S.: 546/146,149; 514/307				
Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched				
NONE				
Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) CAS ONLINE				
C. DOCUMENTS CONSIDERED TO BE RELEVANT				
Category *				Relevant to claim No.
Α	Gunnarsson et al. Designing Small, Nonsugar Activators of Antithrombin Using  1-15			
A	Hydropathic Interaction Analyses. J. Med. Chem. 2000. Vol. 45, pages 1233-1243.  Gunnarsson et al. Interaction of Designed Sulfated Falvanoids with Antithrombin: Lessons on the Design of Organic Activatiors. J. Med. Chem. 2002. Vol. 45. pages 4460-4470.			1-15
			·	
Further	documents are listed in the continuation of Box C.		See patent family annex.	
• s	pecial categories of cited documents:	*T *	later document published after the inte	
"A" document defining the general state of the art which is not considered to be of particular relevance			date and not in conflict with the application but cited to undersus principle or theory underlying the invention	
"E" earlier ap	•		document of particular relevance; the considered novel or cannot be consider when the document is taken alone	
"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)		-y -	document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is	
•			combined with one or more other such being obvious to a person skilled in the	documents, such combination
"P" document	document published prior to the international filing date but later than the		"&" document member of the same patent family	
Date of the actual completion of the international search  Date of mailing of the international search				th report
21 October 2004 (21.10.2004)		21 JAN 2005		
Name and mailing address of the ISA/US		Authorized officer White 1 has		
Mail Stop PCT, Atm: ISA/US		Zinna Northington Davis		
Commissioner for Patents P.O. Box 1450		Telephone No. 571-272-1600		
Alexandria, Virginia 22313-1450  Facsimile No. (703) 305-3230				
Lacanune Mo	. (102) 203-3430			

Form PCT/ISA/210 (second sheet) (January 2004)